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FILE 'HOME' ENTERED AT 20:39:17 ON 26 OCT 2004

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=> file uspatall
COST IN U.S. DOLLARS
```

FULL ESTIMATED COST

SINCE FILE TOTAL
ENTRY SESSION
0.21 0.21

FILE 'USPATFULL' ENTERED AT 20:39:29 ON 26 OCT 2004
CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'USPAT2' ENTERED AT 20:39:29 ON 26 OCT 2004 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

=>	e	ribes	g/in		
E1			1	RIBES	CAROLYN W/IN
E2			1	RIBES	DELPHINE/IN
Е3			0>	RIBES	G/IN
E4			2	RIBES	GERARD/IN
E5			6	RIBES	HERVE/IN
E6			1	RIBES	JOAN MARIA MAS/IN
E7			2	RIBES	LAURENT/IN
E8		*	2	RIBES	MAURICIO/IN

```
RIBES MICHEL/IN
Ε9
             3
E10
             2
                   RIBESSE JACQUES/IN
                   RIBESSE JACQUES MAX/IN
E11
             1
                   RIBET JEAN PAUL/IN
E12
             1
=> s e4
             2 "RIBES GERARD"/IN
L1
=> d 1-2
     ANSWER 1 OF 2 USPATFULL on STN
L1
       Peferences
       2004:38223 USPATFULL
AN
       Pharmaceutical or dietary composition containing a vegetable oil, in
ΤI
       particular olive oil and sitosterol
       Maurel, Jean-Claude, Castries, FRANCE
IN
       Cros, Gerard, Montpellier, GERMANY, FEDERAL REPUBLIC OF
       Ribes, Gerard, Montpellier, FRANCE
       US 2004028759
                      A1
                                20040212
_{PI}
       US 2003-415092
AΙ
                          Α1
                                20030908 (10)
       WO 2001-EP12757
                                20011026
       GB 2000-26609
                           20001031
PRAI
       Utility
DT
       APPLICATION
FS
LN.CNT 479
       INCLM: 424/769.000
INCL
       NCLM: 424/769.000
NCL
       [7]
IC
       ICM: A61K035-78
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
     ANSWER 2 OF 2 USPATFULL on STN
       95:105869 USPATFULL
ΑN
TI
       Treatment of non-insulin-dependent diabetes
IN
       Sauvaire, Yves, Montferrier sur Lez, France
       Ribes, Gerard, Montpellier, France
PA
       Laboratories Monal, France (non-U.S. corporation)
       US 5470879
                                19951128
PΙ
ĀΙ
       US 1994-364079
                               19941223 (8)
       Division of Ser. No. US 1993-113951, filed on 31 Aug 1993, now abandoned
RLI
       FR 1992-10644
                            19920907
       Utility
       Granted
LN.CNT 431
INCL
       INCLM: 514/561.000
NCL
       NCLM: 514/561.000
IC
       ICM: A61K035-78
       514/561
EXF
=> e taouis m/in
E1
             1
                   TAOUFIK MOSTAPHA/IN
E2
                   TAOUIL TONY F/IN
             0 --> TAOUIS M/IN
```

E5

E6

E7

E8

Ε9

1

1

1

1

7

3

TAOUK MICHAEL Y/IN

TAOYAMA MINORU/IN

TAP GEERT E/IN

TAP HENRY H/IN

TAPA BARNABAS/IN

TAOULI ADBERRAHIM/IN

```
1
                    TAPADAR INDRANIL B/IN
E10
E11
                    TAPADAR INDRANIL BOB/IN
             1
E12
             1
                    TAPAI ANTAL/IN
=> e petit r/in
                  PETIT PIERRE H F/IN
E1
              1
E2
              1
                    PETIT PIERRE MARIE/IN
E3
              0
               --> PETIT R/IN
E4
             1
                    PETIT RAYMOND/IN
E5
             2
                    PETIT RAYMOND C/IN
E6
             1
                    PETIT REGIS/IN
             28
                    PETIT ROBERT/IN
E7
             7
                    PETIT ROBERT G/IN
E8
             2
                    PETIT ROBERT G II/IN
Ε9
E10
             1
                    PETIT ROBERTS CORAL A/IN
E11
             3
                    PETIT ROGER/IN
                    PETIT ROGER M/IN
E12
             1
=>
=> e
             3
                    PETIT ROLAND/IN
E13
                    PETIT SAMUEL/IN
E14
             3
E15
            10
                    PETIT SERGE/IN
                    PETIT STEPHANE/IN
E16
             9
                    PETIT STEVEN HAROLD/IN
E17
             1
                    PETIT STEVEN M/IN
E18
             1
                    PETIT THIERRY/IN
E19
             1
                    PETIT THOMAS J/IN
E20
             4
                    PETIT TODD MICHEAL/IN
             1
E21
E22
             1
                    PETIT VINCENT/IN
E23
            12
                    PETIT WILLIAM A/IN
E24
             3
                    PETIT XAVIER/IN
=> s e11
L2
              3 "PETIT ROGER"/IN
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=> d 1-3

L2 ANSWER 1 OF 3 USPATFULL on STN

```
ΑN
       2002:8409 USPATFULL
ΤI
       Portable electronic appliance provided with an integrated radio
       communication devivce having an antenna for transmitting and/or
       receiving electromagnetic waves
IN
       Martinez, Eric, Sartrouville, FRANCE
       Petit, Roger, Sartrouville, FRANCE
PA
       ALCATEL (non-U.S. corporation)
       US 2002004409
PI
                          A1
                                20020110
       US 2001-897139
ΑI
                          A1
                                20010703 (9)
       FR 2000-8965
                           20000710
PRAI
       Utility
DT
FS
       APPLICATION
LN.CNT 294
       INCLM: 455/550.000
INCL
       INCLS: 455/090.000
NCL
       NCLM:
              455/550.000
       NCLS:
              455/090.000
IC
       [7]
       ICM: H04M001-00
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L2 ANSWER 2 OF 3 USPATFULL on STN



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1999:70403 USPATFULL
AΝ
       Identity card reader device
ΤI
       Savalle, Patrick, Rueil Malmaison, France
IN
       Petit, Roger, Sartrouville, France
       Alcatel Mobile Phones, Paris, France (non-U.S. corporation)
PA
                                 19990622
ΡI
       US 5915016
       US 1996-768396
                                 19961218 (8)
AΙ
       FR 1995-15054
                             19951219
PRAI
\overline{\mathrm{DT}}
       Utility
FS
       Granted
LN.CNT 135
       INCLM: 379/433.000
INCL
       INCLS: 235/441.000; 235/492.000
       NCLM: 379/433.090
NCL
       NCLS: 235/441.000; 235/492.000
IC
       [6]
       ICM: H04M001-00
       235/441; 379/433; 379/369
EXF
     ANSWER 3 OF 3 USPATFULL on STN
L2
          an elejek
        Pelerences
       91:11991 USPATFULL
ΑN
       Mouthpiece for wind instrument, and corresponding ligature and
TΙ
       mouthpiece cover
       Petit, Roger, 27, boulevard de 1'Est, 93340 Le Raincy, France
IN
       US 4991483
                                 19910212
ΡI
       US 1989-454942
                                 19891222 (7)
AΙ
       FR 1988-17117
                             19881223
PRAI
\overline{\mathrm{DT}}
       Utility
       Granted
FS
LN.CNT 413
       INCLM: 084/383.000R
INCL
NCL
       NCLM: 084/383.000R
IC
       [5]
       ICM: G10D009-02
       084/383R
EXF
=> e broca c/in
E1
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                    BROBYN SUSAN E/IN
              1
                    BROC GUILLAUME/IN
E2
              0 --> BROCA C/IN
E.3
                    BROCARD DIDIER/IN
F. 4
              1
E.5
              3
                    BROCARD EMMANUEL/IN
F.6
             1
                    BROCARD FABRICE/IN
                    BROCARD FRANCOIS/IN
E7
             1
                    BROCARD HUGUETTE/IN
E8
             1
             1
                    BROCARD JACQUES/IN
E9
E10
             1
                    BROCARD JACQUES BERTRAND/IN
             23
                    BROCARD JEAN MARIE/IN
E11
                    BROCART ALAIN/IN
E12
=> e broca/in
              1
                    BROBYN SUSAN E/IN
E1
E2
              1
                    BROC GUILLAUME/IN
Е3
              0 --> BROCA/IN
F. 4
              1
                    BROCARD DIDIER/IN
              3
                    BROCARD EMMANUEL/IN
E.5
                    BROCARD FABRICE/IN
E6
             1
E7
             1
                    BROCARD FRANCOIS/IN
             1
                    BROCARD HUGUETTE/IN
E8
             1
                    BROCARD JACQUES/IN
E9
              1
                    BROCARD JACQUES BERTRAND/IN
E10
```

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23
                    BROCARD JEAN MARIE/IN
E.11
E12
             1
                    BROCART ALAIN/IN
=> e sauvaire/in
                    SAUVAIGO SYLVIE/IN
              4
E1
                    SAUVAIN ROGER/IN
E2
              1
              0 --> SAUVAIRE/IN
E3
E4
             2
                    SAUVAIRE YVES/IN
                    SAUVAJOL RENE/IN
E5
             1
                    SAUVAN GERARD/IN
             2
E6
             2
                    SAUVAN JACQUES/IN
E7
             4
                    SAUVAN JACQUES LOUIS/IN
E8
F. 9
             1
                    SAUVAN MARCEL/IN
             5
                    SAUVANET MAURICE/IN
E10
E11
             1
                    SAUVANET MAURICE M/IN
E12
              2
                    SAUVANT MOYNOT VALERIE/IN
=> s e4
              2 "SAUVAIRE YVES"/IN
L3
=> d 1-2
     ANSWER 1 OF 2 USPATFULL on STN
         95:105869 USPATFULL
ΑN
TΙ
       Treatment of non-insulin-dependent diabetes
       Sauvaire, Yves, Montferrier sur Lez, France
IN
       Ribes, Gerard, Montpellier, France
       Laboratories Monal, France (non-U.S. corporation)
PA
                                19951128
       US 5470879
ΡĪ
       US 1994-364079
\overline{\mathsf{AI}}
                                19941223 (8)
\overline{\mathtt{RLI}}
       Division of Ser. No. US 1993-113951, filed on 31 Aug 1993, now abandoned
       FR 1992-10644
                            19920907
PRAI
DT
       Utility
       Granted
FS
LN.CNT 431
       INCLM: 514/561.000
INCL
NCL
       NCLM: 514/561.000
IC
       [6]
       ICM: A61K035-78
       514/561
EXF
L3
     ANSWER 2 OF 2 USPATFULL on STN
        references
       92:9044 USPATFULL
AN
       Process for the accelerated ageing and treatment of iris rhizomes
TI
IN
       Baccou, Jean-Claude, Montpellier, France
       Bessiere, Jean-Marie, St. Clement, France
       Boisseau, Patrick, Provence, France
       Faugeras, Pierre, Pour St. Espris, France
       Jouy, Nicholas, Juvignac, France
       Peyrot, Elysabette, Saint Cyr Au Mo, France
       Sauvaire, Yves, Montferrier Sur, France
       Commissariat a l'Energie Atomique, both of, France (non-U.S. government)
PA.
       Universite des Sciences et Techniques du Languedoc, both of, France
       (non-U.S. corporation)
                                19920204
PΙ
       US 5085994
       US 1990-604572
ΑI
                                19901026 (7)
       FR 1989-14042
                            19891026
PRAI
       Utility
DT
       Granted
FS
LN.CNT 244
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INCL INCLM: 435/148.000 INCLS: 435/173.000

NCL NCLM: 435/148.000

NCLS: 435/173.800

IC [5]

ICM: C12P007-26 ICS: C12N013-00

EXF 435/148; 435/173

=> file merck

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
22.64
22.85

FILE 'MRCK' ENTERED AT 20:46:32 ON 26 OCT 2004 COPYRIGHT (C) 2004 Merck & Co., Inc., Whitehouse Station, New Jersey, USA. All Rights

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=> e hydroxyisoleucine

E1	1	HYDROXYISOEUGENOL/BI
E2	1	HYDROXYISOINDOLINE/BI
E3	0>	HYDROXYISOLEUCINE/BI
E4	1	HYDROXYISONITROSOPROPIOPHENONE/BI
E5	1	HYDROXYISOPHTHALIC/BI
E6	1	HYDROXYISOPROPYL/BI
E7	2	HYDROXYISOPROPYLAMINE/BI
E8	1	HYDROXYISOVALERIC/BI
E9	1	HYDROXYISOXAZOLE/BI
E10	1	HYDROXYKAUR/BI
E11 .	45	HYDROXYL/BI
E12	23	HYDROXYLAMINE/BI

=> e 4-hydroxyisoleucine

1		3ZN/BI
5786		4/BI
0	>	4-HYDROXYISOLEUCINE/BI
800		40/BI
229		400/BI
68		4000/BI
5		40000/BI
1		400000/BI
1		40002/BI
4		4001/BI
1		4001323/BI
1		40014/BI
	0 800 229 68 5 1 1	0> 800 229 68 5 1

=> file medline

COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	0.60	23.45

FILE 'MEDLINE' ENTERED AT 20:47:42 ON 26 OCT 2004

FILE LAST UPDATED: 26 OCT 2004 (20041026/UP). FILE COVERS 1951 TO DATE.

On February 29, 2004, the 2004 MeSH terms were loaded. See $\underline{\text{HELP RLOAD}}$ for details. OLDMEDLINE now back to 1951.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2004 vocabulary. See http://www.nlm.nih.gov/mesh/ and http://www.nlm.nih.gov/pubs/techbull/nd03/nd03_mesh.html for a description of changes.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 4-hydroxyisoleucine

2051551 4

15 HYDROXYISOLEUCINE

L4 10 4-HYDROXYISOLEUCINE

(4(W) HYDROXYISOLEUCINE)

=> d 1-10

L4 ANSWER 1 OF 10 MEDLINE on STN

Full sping Text References

AN 2004443395 MEDLINE

- DN PubMed ID: 15350674
- TI Genotoxicity testing of a fenugreek extract.
- AU Flammang A M; Cifone M A; Erexson G L; Stankowski L F Jr
- CS Ross Products Division, Abbott Laboratories, 624 Cleveland Ave, Columbus, OH 43215-1724, USA.. ann.flammang@abbott.com
- SO Food and chemical toxicology: an international journal published for the British Industrial Biological Research Association, (2004 Nov) 42 (11) 1769-75.

Journal code: 8207483. ISSN: 0278-6915.

- CY England: United Kingdom
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals
- EM 200410
- ED Entered STN: 20040908

Last Updated on STN: 20041022 Entered Medline: 20041021

L4 ANSWER 2 OF 10 MEDLINE on STN

Full City Text References

- AN 2004404025 MEDLINE
- DN PubMed ID: 15082420
- TI Insulinotropic agent ID-1101 (4-hydroxyisoleucine) activates insulin signaling in rat.
- AU Broca Christophe; Breil Vincent; Cruciani-Guglielmacci Celine; Manteghetti Michele; Rouault Christine; Derouet Michel; Rizkalla Salwa; Pau Bernard; Petit Pierre; Ribes Gerard; Ktorza Alain; Gross Rene; Reach Gerard; Taouis Mohammed
- CS Laboratoire de Pharmacologie, Centre de Pharmacologie et Biotechnologies pour la Sante-Unite Mixte de Recherche 5160 Centre National de la Recherche Scientifique, Faculte de Medecine, 34060 Montpellier, France.. christophe.broca@univ-montp1.fr
- American journal of physiology. Endocrinology and metabolism, (2004 Sep) 287 (3) E463-71.
 - Journal code: 100901226. ISSN: 0193-1849.
- CY United States
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals
- EM 200409
- ED Entered STN: 20040814

Last Updated on STN: 20040911 Entered Medline: 20040910

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ANSWER 3 OF 10
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         2002736689
                    MEDITNE
ΑN
     PubMed ID: 12498629
DN
ΤI
     New legume sources as therapeutic agents.
ΑIJ
     Madar Zecharia; Stark Aliza H
     The Hebrew University of Jerusalem, Faculty of Agricultural, Food and
CS
     Environmental Quality Sciences, Institute of Biochemistry, Food Science
     and Nutrition, P.O. Box 12, Rehovot, 76100, Israel.. madar@agri.huji.ac.il
     British journal of nutrition, (2002 Dec) 88 Suppl 3 S287-92. Ref: 45
SO
     Journal code: 0372547. ISSN: 0007-1145.
     England: United Kingdom
CY
     Journal; Article; (JOURNAL ARTICLE)
DT
     General Review; (REVIEW)
     (REVIEW, TUTORIAL)
     English
LA
FS
     Priority Journals
     200302
EM
     Entered STN: 20021227
ED
     Last Updated on STN: 20030221
     Entered Medline: 20030220
     ANSWER 4 OF 10
                        MEDLINE on STN
L4
       Peterences
     2002093541
                    MEDITNE
AΝ
     PubMed ID: 11740101
DN
     Full stereochemical understanding in a new (2R, 3R, 4R)-4-
TΤ
     hydroxyisoleucine synthesis.
     Rolland M; Kassem T; Rolland V; Martinez J
ΑIJ
     Laboratoire des Aminoacides, Peptides et Proteines, Universite Montpellier
CS
     I et II, Faculte de Pharmacie, UMR CNRS 5810, 15 Avenue Charles Flahault,
     34060 Montpellier CEDEX 2, France.. rolland@colombes.pharma.univ-montp1.fr
     Acta crystallographica. Section C, Crystal structure communications, (2001
SO
     Dec) 57 (Pt 12) 1415-7.
     Journal code: 8305826. ISSN: 0108-2701.
CY
     Denmark
DT
     Journal; Article; (JOURNAL ARTICLE)
     English
LΑ
FS
     PUBMED-NOT-MEDLINE
     200202
EM
     Entered STN: 20020202
ED
     Last Updated on STN: 20020227
     Entered Medline: 20020226
     ANSWER 5 OF 10
                        MEDLINE on STN
L4
     2000469059
                    MEDLINE
ΑN
     PubMed ID: 10944320
DN
     Two key chiral intermediates in a new 4-hydroxyisoleucine synthesis.
TI
     Kassem T; Rolland V; Martinez J; Rolland M
ΑU
     Laboratoire des Aminoacides, Peptides et Proteines, Universite Montpellier
CS
     I et II, Faculte de Pharmacie, UMR CNRS 5810, 15 Avenue Charles Flahault,
     34060 Montpellier CEDEX 2, France.
     Acta crystallographica. Section C, Crystal structure communications, (2000
SO
     Aug) 56 ( Pt 8) 1037-9.
     Journal code: 8305826. ISSN: 0108-2701.
CY
     Denmark
     Journal; Article; (JOURNAL ARTICLE)
DT
     English
LA
FS
     Priority Journals
EM
     200010
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Entered STN: 20001012

Last Updated on STN: 20001012 Entered Medline: 20001005

L4 ANSWER 6 OF 10 MEDLINE on STN

Full Text

- AN 2000175398 MEDLINE
- DN PubMed ID: 10708743
- TI 4-Hydroxyisoleucine: effects of synthetic and natural analogues on insulin secretion.
- AU Broca C; Manteghetti M; Gross R; Baissac Y; Jacob M; Petit P; Sauvaire Y; Ribes G
- CS UMR 9921 du Centre National de la Recherche Scientifique, Montpellier, France.. broca@zeus.sc.univ-montp1.fr
- SO European journal of pharmacology, (2000 Mar 3) 390 (3) 339-45. Journal code: 1254354. ISSN: 0014-2999.
- CY Netherlands
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals
- EM 200005
- ED Entered STN: 20000518

Last Updated on STN: 20000518 Entered Medline: 20000511

L4 ANSWER 7 OF 10 MEDLINE on STN

Full State Text References

- AN 1999447236 MEDLINE
- DN PubMed ID: 10516120
- TI 4-Hydroxyisoleucine: experimental evidence of its insulinotropic and antidiabetic properties.
- AU Broca C; Gross R; Petit P; Sauvaire Y; Manteghetti M; Tournier M; Masiello P; Gomis R; Ribes G
- CS Unite Mixte de Recherche 9921 du Centre National de la Recherche Scientifique, Faculte de Medecine UPRES EA 1677, 34060 Montpellier, France. broca2zeus.sc.univ-montp1.fr.
- SO American journal of physiology, (1999 Oct) 277 (4 Pt 1) E617-23. Journal code: 0370511. ISSN: 0002-9513.
- CY United States
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals
- EM 199911
- ED Entered STN: 20000111

Last Updated on STN: 20000111 Entered Medline: 19991122

L4 ANSWER 8 OF 10 MEDLINE on STN

Full sens Text Sereignes

- AN 1998178529 MEDLINE
- DN PubMed ID: 9519714
- TI 4-Hydroxyisoleucine: a novel amino acid potentiator of insulin secretion.
- AU Sauvaire Y; Petit P; Broca C; Manteghetti M; Baissac Y; Fernandez-Alvarez J; Gross R; Roye M; Leconte A; Gomis R; Ribes G
- CS Laboratoire de Recherche sur les Substances Naturelles Vegetales,
- Universite Montpellier II, France. SO Diabetes, (1998 Feb) 47 (2) 206-10.
 - Journal code: 0372763. ISSN: 0012-1797.
- CY United States
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Abridged Index Medicus Journals; Priority Journals
- EM 199804

ED Entered STN: 19980416

Last Updated on STN: 19980416 Entered Medline: 19980403

L4 ANSWER 9 OF 10 MEDLINE on STN

Full Communication Full Text References

- AN 97194135 MEDLINE
- DN PubMed ID: 9041713
- TI Characterization of a dioxygenase from Trigonella foenum-graecum involved in 4-hydroxyisoleucine biosynthesis.
- AU Haefele C; Bonfils C; Sauvaire Y
- CS Laboratoire de Recherche sur les Substances Naturelles Vegetales, UPR ES 1677, CP 024, Universite Montpellier II, France.
- SO Phytochemistry, (1997 Feb) 44 (4) 563-6. Journal code: 0151434. ISSN: 0031-9422.
- CY United States
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Biotechnology
- EM 199703
- ED Entered STN: 19970407

Last Updated on STN: 19970407 Entered Medline: 19970324

L4 ANSWER 10 OF 10 MEDLINE on STN

Full Text References

AN 79224186 MEDLINE

- DN PubMed ID: 461558
- TI The detection of isomers of 4-hydroxyisoleucine by the Jeol Amino Acid Analyser and by TLC. The detection of isomers of 4-hydroxyisoleucine by the Jeol Amino Acid Analyser and by TLC.
- AU Hardman R; Abu-Al-Futuh I M
- SO Planta medica, (1979 May) 36 (1) 79-84.
 - Journal code: 0066751. ISSN: 0032-0943.
- CY GERMANY, WEST: Germany, Federal Republic of
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals
- EM 197909
- ED Entered STN: 19900315

Last Updated on STN: 19900315 Entered Medline: 19790925

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L4 ANSWER 8 OF 10 MEDLINE on STN



- AN 1998178529 MEDLINE
- TI 4-Hydroxyisoleucine: a novel amino acid potentiator of insulin secretion.
- AU Sauvaire Y; Petit P; Broca C; Manteghetti M; Baissac Y; Fernandez-Alvarez J; Gross R; Roye M; Leconte A; Gomis R; Ribes G
- SO Diabetes, (1998 Feb) 47 (2) 206-10. Journal code: 0372763. ISSN: 0012-1797.
- TI 4-Hydroxyisoleucine: a novel amino acid potentiator of insulin secretion.
- AB We report the characterization of a new insulinotropic compound,

 4-hydroxyisoleucine. This amino acid has been extracted and purified from fenugreek seeds, which are known in traditional medicine for their antidiabetic properties. 4-Hydroxyisoleucine increases glucose-induced insulin release, in the concentration range of 100 micromol/l to 1 mmol/l, through a direct effect on isolated islets of Langerhans from both rats

and humans. The stimulating effect of 4-hydroxyisoleucine was strictly glucose dependent; indeed, ineffective at low (3 mmol/1) or basal (5 mmol/1) glucose concentrations, the amino acid potentiated. . . In addition, in the isolated perfused rat pancreas, we could show 1) that the pattern of insulin secretion induced by 4-hydroxyisoleucine was biphasic, 2) that this effect occurred in the absence of any change in pancreatic alpha- and delta-cell activity, and 3) that the more glucose concentration was increased, the more insulin response was amplified. Moreover, 4-hydroxyisoleucine did not interact with other agonists of insulin secretion (leucine, arginine, tolbutamide, glyceraldehyde). Therefore, we conclude that 4-hydroxyisoleucine insulinotropic activity might, at least in part, account for fenugreek seeds' antidiabetic properties. This secretagogue may be considered as a. 0 (4-hydroxyisoleucine); 0 (Hypoglycemic Agents); 0 (Plant Extracts) We report the characterization of a new insulinotropic compound, 4-hydroxyisoleucine. This amino acid has been extracted and purified from fenugreek seeds, which are known in traditional medicine for their antidiabetic properties. 4-Hydroxyisoleucine increases glucose-induced insulin release, in the concentration range of 100 micromol/l to 1 mmol/l, through a direct effect on isolated islets of Langerhans from both rats and humans. The stimulating effect of 4-hydroxyisoleucine was strictly glucose dependent; indeed, ineffective at low (3 mmol/1) or basal (5 mmol/1) glucose concentrations, the amino acid potentiated the insulin secretion induced by supranormal (6.6-16.7 mmol/1) concentrations of glucose. In addition, in the isolated perfused rat pancreas, we could show 1) that the pattern of insulin secretion induced by 4-hydroxyisoleucine was biphasic, 2) that this effect occurred in the absence of any change in pancreatic alpha- and delta-cell activity, and 3) that the more glucose concentration was increased, the more insulin response was amplified. Moreover, 4-hydroxyisoleucine did not interact with other agonists of insulin secretion (leucine, arginine, tolbutamide, glyceraldehyde). Therefore, we conclude that 4-hydroxyisoleucine insulinotropic activity might, at least in part, account for fenugreek seeds' antidiabetic properties. This secretagogue may be considered as a novel drug with potential interest for the

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treatment of NIDDM.